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4	Orienteering combines vigorous-intensity exercise with navigation to
5	improve human cognition and increase brain-derived neurotrophic
6	factor
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## 25 Abstract

- 26 Exercise enhances aspects of human cognition, but its intensity may matter. Recent animal
- 27 research suggests that vigorous exercise, which releases greater amounts of lactate, activates
- 28 more brain-derived neurotrophic factor (BDNF) in the hippocampus and, thus, may be optimal

- 71 exponentially beyond the lactate threshold of ~ 4mmol/L of lactate in untrained adults [11,12].
- Although lactate has historically and erroneously been considered an inert metabolic waste [13],
- recent evidence points to its importance as both a fuel source [14] and an activator of BDNF

[15–18] with rapid effects. Mere m

# Participants

Effects	of	orien	teering	on	lactate.	<b>BDNF</b>	and	cognition
					,			

163	exercise. The total activity amount was determined by multiplying the average length of an
164	exercise session by the average number of active days (minutes/week).
165	The Navigational Strategy Questionnaire (NSQ) was used to assess participants' baseline
166	navigational tendencies [47]. Using a 5-point Likert scale, participants rated 44 items
167	corresponding to three different navigational strategies: allocentric spatial processing, egocentric
168	spatial processing, and procedural processing. For each strategy, an average score was
169	calculated.
170	Baseline autobiographical memory was assessed using the Survey of Autobiographical
171	Memory (SAM; [48]. In the SAM, subjective memory is evaluated across 26 items which are
172	answered using a 5-point Likert scale. Each item is weighted and summed to obtain an average
173	for four memory domains including episodic, spatial, semantic, and future memory. In this study,
174	we examined episodic and spatial memory specifically.
175	Lab-based baseline measurements

In the lab, before the intervention, the participant's height (centimetres), weight

- 186 (intensity) + HR<sub>Rest</sub>. For the moderate-intensity orienteering group, the exercise intensity range
- 187 was calculated as 40-50% of HRR, and 80-85% of HRR was used for the vigorous-intensity
- 188 orienteering and vigorous-intensity exercise groups.
- 189 Estimates of VO<sub>2</sub> peak were calculated using the *WorldFitnessLevel.org* website [50].
- 190 Participants were asked to respond to the website questions as accurately as possible and input
- 191 their anthropometric and HR measurements.

#### 192 Intervention measures of exercise intensity

193During the intervention, HR, ratings of perceived exertion (RPE) and lactate were

194 recorded at the middle and end of the intervention course and 10 minutes post-intervention. The

195 highest of these values was analyzed. Heart rate was recorded using the Polar HR-10 monitor.

196 RPE was captured using the Borg 1-20 Scale [51]. Lactate was measured from a sample of whole

197 blood obtained from the fingertip using the Lactate Plus portable analyzer (Nova Biomedical,

198 Waltham, MA).

#### 199 **Pre- and post-intervention measurements**

Before the intervention, the cognitive testing was completed before obtaining a serum sample for BDNF. Following the intervention, the blood sample was collected within 10 minutes of finishing the intervention course and was followed by cognitive testing.

For BDNF, three-hour fasted samples of venous blood were obtained from a vein in the
antecubital fossa. Samples were collected into BD Vacutainer SST tubes (BD, Franklin Lakes,
NJ), chilled on ice, allowed to clot for a minimum of 45 minutes following sample collection and
then centrifuged at 1000 x g for 15 minutes at 4°C. For all samples, 300µL of supernatant was
collected to obtain serum, aliquoted into microtubes, and stored immediately at -20°C until

208 analysis. The concentration of serum BDNF was quantified using a sandwich Biosensis Mature

209	BDNF Rapid <sup>TM</sup> ELISA Kit (Biosensis Pty Ltd, Thebarton, Australia). Samples were diluted
210	100x and were run in duplicate. Using a BioTek SynergyMx spectrophotometer, absorbance was
210	Toox and were full in duplicate. Using a biorek Synergywix specifophotometer, absorbance was
211	measured at 450 nm and analyzed using Gen 5 1.11 Software (BioTek Instruments Inc.,
212	Winooski, VT). Select samples whose concentration fell above the standard curve of the
213	preliminary analysis were re-analyzed using a 125x dilution and the same protocol.
214	Memory was tested in two ways. First, memory was tested using Kirwan and Stark's
215	Mnemonic Similarity Task [52–54], a modified object recognition task that places a large
216	emphasis on high-interference memory and hippocampal function. The Mnemonic Similarity
217	Task begins with a study phase in which participants are shown a series of images of 60
218	everyday objects displayed on the screen for two seconds and must classify whether the image is
219	an 'indoor' or an 'outdoor' item. This is immediately followed by a test phase, in which
220	participants are shown 20 'repeat' images (correct response = "old"), 20 'lure' images that are
221	highly similar but not identical to a previous image (correct response = "similar"), and 20
222	completely new, 'foil,' images (correct response = "new") and asked to classify them. The
223	Mnemonic Similarity Task has two measures which provide a valuable distinction between
224	hippocampal-dependent high-interference memory and recognition memory. The "lure
225	discrimination index" is a measure of high-interference memory, calculated as [ $p$ ("Similar"
226	Lure image) – $p$ ("Similar"   Foil image)] × 100, and reflects one's ability to correctly classify
227	'lure' items as "similar". High-interference memory relies on the ability to remember specific
228	details during encoding [55], which is dependent on the function of the hippocampus and is
229	associated with hippocampal neurogenesis [54]. The second measure of the Mnemonic Similarity
230	Task is general "recognition memory", defined as the ability to correctly label a 'repeat' image
231	as "old," [ $p$ ("Old"   Repeat image) – $p$ ("Old"   Foil image)] × 100. Recognition memory does

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not require

- learned how to read their HR on the Polar Pacer Pro GPS watch and maintain their pace so that
- their HR remained in the target range. Participants in the orienteering groups (both moderate and
- 257 vigorous intensity) were taught how to use the orienteering map legend, orient their map, use the map to plan a route and locate checkpoints, and re-

- their navigational decisions. In contrast, those in the exercise only group exercised at a vigorous
- 279 intensity (80-85% of HRR) but did not engage in orienteering. Instead, a member of the research
- team led the participant along the most efficient route.
- 281 All participants were responsible for tracking their HR at each checkpoint and were
- 282 instructed to adapt the pace or pause until their HR returned to the target zone for a maximum of one minute. At the midpoint and finish

300	To test for potential covariates, a one-way analysis of variance (ANOVA) was used to
301	assess group differences in all demographic variables, weekly physical activity, spatial
302	navigation tendencies and autobiographical memory, as well as pre-intervention differences in
303	BDNF and cognition. To ensure that our intervention was adequate in reaching the desired
304	exercise intensity, one-way ANOVA tests were computed for peak HR, peak RPE and peak
305	blood lactate between groups. For blood lactate, a Kruskal-Wallis Means Ranks Test was used to
	confirm that the proportions of those above or below the lactate threshold of 4mmol/>f,()] 488qQq0.00000912 (

- 323 test efficiency). Then, we performed an exploratory analysis using Spearman's correlation to
- 324 evaluate the relationship between the composite cognition score with peak lactate and percent
- 325 change in BDNF. Finally, we conducted a partial Spearman's correlation to determine whether
- 326 the association between composite cognition score and peak lactate was diminished after
- 327 controlling for the percent change in BDNF.

#### Secondary outcome

- 346 and memory. However, pre-intervention BDNF levels were higher for the moderate orienteering
- 347 group than the vigorous orienteering or vigorous exercise groups (p < .001) (Table 2). Univariate
- 348 ANOVA tests confirmed no other baseline differences between groups (Table 1).
- 349
- 350 **Table 1. Descriptive Statistics Between Intervention Groups**

Efficiency	$26.12\pm10.32$	$23.28\pm 6.85$	$24.60\pm5.89$
Pre	$29.45\pm7.88$	$29.42\pm6.45$	$29.92 \pm 4.52$

Post

## Groton Maze Test Efficiency

Pre

Post

437

## 438 **Peak lactate, BDNF and cognitive function**

In this exploratory analysis, the Spearman's

496	Pre				
497 498 499	4. Groton Maze Learning Efficiency Pre	29	.20	.08	
500 501	5. Groton Maze				
502	* = p < .05, *** = p < .001.				
503					

#### 504 **Discussion**

505 The present study was the first to examine the effects of an acute bout of orienteering 506 versus exercise on cognition in a sample of healthy young adults who were recreationally active 507 but unfamiliar with orienteering. The results revealed a strong effect of exercise intensity such 508 that the vigorous-intensity interventions in the form of either running or orienteering elicited 509 greater increases in lactate, BDNF and memory than the moderate-intensity intervention. 510 Additionally, vigorous orienteering improved spatial learning and memory more than vigorous 511 running, suggesting an additional benefit of simultaneous training. 512 This study demonstrates a link between lactate, BDNF and cognition in humans. A novel 513 and important finding is that the higher peak lactate induced by our vigorous exercise 514 interventions was associated with greater percent increases in BDNF and better memory than our 515 moderate-intensity intervention, lending support for the hypothesis that lactate mediates muscle-516 to-brain signalling [10,15,16,19]. Cognition was also significantly related to peak levels of 517 lactate obtained during exercise. Interestingly, when controlling for BDNF, the relationship 518 between cognition and lactate was no longer significant. We hypothesize that BDNF may partly 519 underlie the effects of lactate on cognition, however, further work is needed to understand how 520 exercise-induced lactate impacts cognition through and beyond its effects on BDNF 521 [10,15,16,19].

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522 On top of vigorous-intensity effects, running while navigating conferred additional 523 benefits on our measure of spatial cognition. Spatial learning and memory were tested using the 524 Groton Maze Learning Test, which is a close 2D analog to the 3D wayfinding of orienteering. 525 Although all groups increased in spatial learning efficiency, the vigorous orienteering group 526 improved the most and was the only group to improve in spatial memory after a delay. It is 527 important to consider why. One reason relates to the specific cognitive processes tested. During 528 the Groton Maze Learning Test, participants had to recall the maze route immediately and after a 529 10-minute delay, requiring skills that are highly dependent on the hippocampus, a brain region 530 that is responsive to intervention-induced plasticity [60]. A second reason why orienteering may 531 preferentially benefit spatial cognition relates to its overlap in cognitive processes engaged by 532 the task. In general, cognitive training effects tend to transfer more readily to "near-transfer" 533 tasks, i.e., tasks that closely resemble the cognitive demands of the training protocol, than "far-534 transfer" tasks, i.e., tasks that depend on more disparate cognitive processes [61,62]. In the case 535 of orienteering, spatial cognition would classify as a near-transfer task and based on this 536 framework, would be expected to benefit the most.

537 In contrast, the high-interference memory task would be considered a far-transfer task 538 and, by the same logic, would be less likely to show additive effects, as was observed. Instead, 539 high-interference memory (lure discrimination index) improved to a similar extent for both 540 vigorous exercise and orienteering groups, suggesting that this aspect of cognition is more 541 sensitive to the acute effects of exercise intensity than the combined effects of the exercise-542 cognitive training that is experienced during an acute bout of orienteering. Although the effects 543 of vigorous exercise on high-interference memory were expected and consistent with prior work [9.56.57.63], we were surprised to observe a decrement in high-interference memory 544

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545	performance following moderate-intensity orienteering. This may be related to the amount of
546	exercise-induced BDNF, which is less after moderate intensities compared to vigorous [6-8].
547	Indeed, those who orienteered at moderate intensity produced less BDNF than those who
548	orienteered at a vigorous intensity, and this may have reduced their neurogenic support,
549	rendering substrate-dependent memory benefits unobtainable.
550	The difference in BDNF levels between moderate and vigorous intensity orienteering
551	may also help to explain why expert taxi drivers experience a trade-off that augments their
552	posterior (primarily relating to spatial processing) hippocampus at the cost of their anterior
553	(mainly involved in episodic memory) hippocampus [39]. Taxi drivers are sedentary while
554	driving, which is in stark contrast to expert orienteers who perform their sport at a rapid running
555	speed [29]. The lack of vigorous movement during navigation may be why we see evidence for a
556	trade-off in expert taxi drivers but not in expert orienteers. Regardless of the mechanism, we
	found that engaging in vigorous-intensity exercise while

568	that BDNF is thought to respond to energetic challenges [65], and in our sample of recreationally
569	active younger adults, the additional challenge of running while navigating may not have been
570	enough of an acute energetic demand. This may be especially true given the wayfinding task was
571	short (only ~12 minutes) and across a familiar terrain. Future work should examine the
572	potentially additive effects of orienteering versus running on BDNF using longer and less
573	familiar routes. Additionally, there is evidence that females have lower BDNF responsivity to
574	acute exercise [66] and lower lactate responses at the same relative exercise intensity compared
575	to males [67]. These potential sex-based differences in lactate-induced BDNF activation may be
576	at play with our predominately female sample (65%) and should be followed up in future work.
577	Despite our participants' familiarity with the campus on which the orienteering course
578	was set, both the moderate and vigorous intensity orienteering groups travelled significantly
579	farther and, by extension, made more errors than the most efficient route. Interestingly, the
580	distance travelled while orienteering was associated with several of our baseline measures.
581	Notably, those who travelled shorter distances (i.e., made fewer errors) had better spatial
582	memory at baseline, as revealed by both self-report and task performance, which reaffirms the
583	existence of overlapping cognitive processes engaged between navigation and spatial memory
584	[30]. Also, those who travelled shorter distances reported greater reliance on egocentric spatial
585	navigation. Allocentric spatial navigation was not as strongly related to course distance travelled,
586	which was surprising given that allocentric spatial processing, like egocentric spatial processing
587	and spatial memory, have been previously associated with expertise in the sport of orienteering
588	[27]. The weak association between allocentric spatial processing and navigational efficiency

acknowledge that navigational tendencies may differ between familiar and unfamiliar terrains
[30]. For example, participants could identify campus buildings by their names and then navigate
based on previously learned routes rather than utilize allocentric spatial navigation. It will be
important for future work to examine the orienteering interventions across novel terrains over a
variety of course difficulties.

596 Moreover, overreliance on GPS devices may be a factor because it minimizes active 597 navigation and the practice of allocentric navigation in the case of "use it or lose it" [36]. GPS 598 may be used more commonly by those with little experience in orienteering, as allocentric 599 navigation may require more practice to be developed [30]. Unfortunately, we did not capture 600 GPS use, but we would recommend this be done in future studies. Furthermore, prior research 601 suggests that females may rely less on allocentric navigation than males [68], and our sample 602 was predominantly female. Although we did not power our sample size to examine sex 603 differences, it is recommended that future research do so. Another reason why we failed to 604 observe a strong association between allocentric spatial processing and navigational efficiency is 605 speculative but worth noting; this study was conducted in North America where orienteering 606 awareness and practice is relatively limited compared to Nordic countries where orienteering is 607 embedded into the school curricula and local cultural activities [69]. This fact should be 608 considered when comparing studies from different countries.

609

### 610 **Conclusion**

This study demonstrates the effect of vigorous exercise on lactate, BDNF and
hippocampal-dependent memory. It also reveals that orienteering may outperform exercise in
improving spatial memory when done at a vigorous intensity. Together, this study establishes the

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